| FORM PTO-1390                                                                                                                                                                                                                                                                                                                                                                                                  | ATTORNEY DOCKET NUMBER                                                                                      |  |  |  |  |  |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------|--|--|--|--|--|
| (REV 12-2001)                                                                                                                                                                                                                                                                                                                                                                                                  | 4439-4047                                                                                                   |  |  |  |  |  |
| TRANSMITTAL LETTER TO THE UNI'<br>DESIGNATED/ELECTED OFFICE (I                                                                                                                                                                                                                                                                                                                                                 |                                                                                                             |  |  |  |  |  |
| CONCERNING A FILING UNDER 35                                                                                                                                                                                                                                                                                                                                                                                   |                                                                                                             |  |  |  |  |  |
| INTERNATIONAL APPLICATION INTERNATION PCT/JP2005/006510 INTERNATION 01 April 2005 (0                                                                                                                                                                                                                                                                                                                           | NAL FILING DATE EARLIEST PRIORITY DATE CLAIMED                                                              |  |  |  |  |  |
| TITLE OF INVENTION                                                                                                                                                                                                                                                                                                                                                                                             |                                                                                                             |  |  |  |  |  |
| CARDIOINHIBITORY/ANTIHYPERTENSIVE NOVEL ENDOGENOUS PHYSIOLOGICALLY ACTIVE PEPTIDE APPLICANTS FOR DO/EO/US                                                                                                                                                                                                                                                                                                      |                                                                                                             |  |  |  |  |  |
| Masayoshi SHICHIRI                                                                                                                                                                                                                                                                                                                                                                                             |                                                                                                             |  |  |  |  |  |
| Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:                                                                                                                                                                                                                                                                                |                                                                                                             |  |  |  |  |  |
| 1. A This is FIRST submission of items concerning a filing under 35 U.S.C. 371.                                                                                                                                                                                                                                                                                                                                |                                                                                                             |  |  |  |  |  |
| 2. This is SECOND or SUBSEQUENT submission of items concerning a filing under 35 U.S.C. 371.                                                                                                                                                                                                                                                                                                                   |                                                                                                             |  |  |  |  |  |
| 3. This express request to begin national examination procedures (35 U.S.C. 371(f)).                                                                                                                                                                                                                                                                                                                           |                                                                                                             |  |  |  |  |  |
| 4.   The US has been elected by the expiration of 19 months from the priority date (Article 31).                                                                                                                                                                                                                                                                                                               |                                                                                                             |  |  |  |  |  |
| 5. A copy of the International Application as filed (35 U.S.C. 371(c)(2))                                                                                                                                                                                                                                                                                                                                      |                                                                                                             |  |  |  |  |  |
| a. ☐ is attached hereto (required only if not communicated by the International Bureau). b. ☑ has been communicated by the International Bureau. c. ☐ is not required, as the applications was filled in the United States Receiving Office (RO/US).                                                                                                                                                           |                                                                                                             |  |  |  |  |  |
| 6. An English translation of the International application as filed (35 U.S.C. 371(c)(2)).                                                                                                                                                                                                                                                                                                                     |                                                                                                             |  |  |  |  |  |
| a. \( \) is attached hereto (\) b. \( \) has been previously submitted under 35 U.S.C. 154(d)(4).                                                                                                                                                                                                                                                                                                              |                                                                                                             |  |  |  |  |  |
| 7. Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))                                                                                                                                                                                                                                                                                                        |                                                                                                             |  |  |  |  |  |
| a. ac statched factor (required only if not communicated by the International Burcas). b. laste been communicated by the International Burcas. c. laste not been made, however, the time limit for making such amendments has NOT expired. d. lo have not been made and will not be made.                                                                                                                      |                                                                                                             |  |  |  |  |  |
| 8.                                                                                                                                                                                                                                                                                                                                                                                                             |                                                                                                             |  |  |  |  |  |
| 9. An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).                                                                                                                                                                                                                                                                                                                                            |                                                                                                             |  |  |  |  |  |
| <ol> <li>An English translation of the annexes to the International Preliminary Examination Report under PCT Article 36<br/>(35 U.S.C. 371(e)(5)).</li> </ol>                                                                                                                                                                                                                                                  |                                                                                                             |  |  |  |  |  |
| Items 11. to 20. below concern document(s) or information in                                                                                                                                                                                                                                                                                                                                                   | icluded.                                                                                                    |  |  |  |  |  |
| 11. An Information Disclosure Statement under 37 CFR 1.97 and 1.98, Form PTO-1449, with a copy ofcited references.                                                                                                                                                                                                                                                                                             |                                                                                                             |  |  |  |  |  |
| 12. An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.                                                                                                                                                                                                                                                                                          |                                                                                                             |  |  |  |  |  |
| 13. A FIRST preliminary amendment – pgs.                                                                                                                                                                                                                                                                                                                                                                       |                                                                                                             |  |  |  |  |  |
| 4. A SECOND or SUBSEQUENT preliminary amendment.                                                                                                                                                                                                                                                                                                                                                               |                                                                                                             |  |  |  |  |  |
| 15. English Translation of: Specification (29 pgs.), Claim                                                                                                                                                                                                                                                                                                                                                     | English Translation of: Specification (29 pgs.), Claims (4 pgs.), Abstract (1 pg.), and Figs. 1-2 (1 sheet) |  |  |  |  |  |
| 16. Translation of Deposit of MICROORGANISMS:                                                                                                                                                                                                                                                                                                                                                                  |                                                                                                             |  |  |  |  |  |
| 17.   A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 35 U.S.C. 1.821-1.825 will be furnished today with a Communication by Express Mail → Express Mail No. EY 825780185 US. The paper copy of the sequence listing (5 pg.) and the Statement Under 37 CFR. § 1.82(2) or § 1.825(2) are attacked between one with the computer-cadeble form for the sequence listing. |                                                                                                             |  |  |  |  |  |
| A second copy of the published international application under 35 U.S.C. 154(d)(4).                                                                                                                                                                                                                                                                                                                            |                                                                                                             |  |  |  |  |  |
| A second copy of the English translation of the international application under 35 U.S.C. 154(d)(4).                                                                                                                                                                                                                                                                                                           |                                                                                                             |  |  |  |  |  |
| 20. Other items or Information: Copy of the first page of the published PCT application – publics                                                                                                                                                                                                                                                                                                              | ation number WO 2005/095609, published on October 13, 2005.                                                 |  |  |  |  |  |

| U.S. APPLICATION NO. (if                                                                                                                                                                                                                | known, see 37 C.F.R. 1.51 | INTERNATIONAL APPL | INTERNATIONAL APPLICATION NO. |                           | ATTORNEY'S DOCKET NO. |  |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------|--------------------|-------------------------------|---------------------------|-----------------------|--|
| TBA                                                                                                                                                                                                                                     | 2000                      | PCT/JP2005/006510  |                               | 4439-4047                 |                       |  |
| 21.                                                                                                                                                                                                                                     |                           |                    | CALCULATIONS                  | PTO USE ONLY              |                       |  |
| BASIC NATIONAL FEE (37 CFR 1.492 (a) (1) - (5) ):                                                                                                                                                                                       |                           |                    |                               |                           |                       |  |
| Neither international preliminary examination fee (37 CFR 1.482)<br>nor international search fee (37 CFR 1.445(a)(2) paid to USPTO                                                                                                      |                           |                    |                               |                           |                       |  |
| and International Search Report not prepared by the EPO or JPO \$1,080.00                                                                                                                                                               |                           |                    |                               |                           |                       |  |
| the attent                                                                                                                                                                                                                              | v                         | 6 (27 CED 1 492    | ) and anid to                 |                           |                       |  |
| International preliminary examination fee (37 CFR 1.482) not paid to USPTO but International Search Report prepared by the EPO or JPOS920.00                                                                                            |                           |                    |                               |                           |                       |  |
|                                                                                                                                                                                                                                         |                           |                    |                               |                           |                       |  |
| International preliminary examination fee (37 CFR 1.482) not paid to USPTO but international search fee (37 CFR 1.445(a)(2) paid to USPTO                                                                                               |                           |                    |                               |                           |                       |  |
| International preliminary examination for paid to USPTO (37 CFR 1.482)                                                                                                                                                                  |                           |                    |                               |                           |                       |  |
| but all claims did not satisfy provisions of PCT Article 33 (1) - (4)\$730.00                                                                                                                                                           |                           |                    | 1                             |                           |                       |  |
| International preliminary examination fee paid to USPTO (37 CFR 1.482)                                                                                                                                                                  |                           |                    |                               | 1                         |                       |  |
| and all claims satisfied provisions of PCT Article 33(1) - (4)\$100.00                                                                                                                                                                  |                           |                    |                               |                           |                       |  |
| ENTER APPROPRIATE BASIC FEE AMOUNT =                                                                                                                                                                                                    |                           |                    | \$ 920                        |                           |                       |  |
| Surcharge of \$130 for furnishing the oath or declaration later than 20 30                                                                                                                                                              |                           |                    | S                             |                           |                       |  |
| months from the earliest claimed priority date (37 CFR 1.492(e)).                                                                                                                                                                       |                           |                    | 3                             |                           |                       |  |
| CLAIMS                                                                                                                                                                                                                                  | NUMBER FILED              | NUMBER EXTRA       | RATE                          |                           |                       |  |
| Total claims                                                                                                                                                                                                                            | 22 - 20 =                 | 2                  | x \$18.00                     | \$ 36                     |                       |  |
| Independent claims                                                                                                                                                                                                                      | 4 - 3 =                   | 1                  | x \$86.00                     | \$ 86                     |                       |  |
| MULTIPLE DEPENDENT CLAIM(S) (if applicable) +\$290.00                                                                                                                                                                                   |                           |                    | +\$290.00                     | \$ 290                    |                       |  |
|                                                                                                                                                                                                                                         |                           |                    | CULATIONS =                   | \$1,332                   |                       |  |
| Reduction of ½ for filing by small entity, if applicable. A Small Entity Statement must also by filed (Note 37 CFR 1.9, 1.27, 1.28).                                                                                                    |                           |                    | S                             |                           |                       |  |
| SUBTOTAL =                                                                                                                                                                                                                              |                           |                    | \$1,332                       |                           |                       |  |
| Processing fee of \$130.00 for furnishing the English translation later than □20 □30 months from the earliest claimed priority date (37 CFR 1.492(f)). +                                                                                |                           |                    | S                             |                           |                       |  |
| TOTAL NATIONAL FEE =                                                                                                                                                                                                                    |                           |                    | \$1,332                       |                           |                       |  |
| Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property +                                                                |                           |                    | \$                            |                           |                       |  |
| TOTAL FEES ENCLOSED                                                                                                                                                                                                                     |                           |                    | \$1,332                       |                           |                       |  |
|                                                                                                                                                                                                                                         |                           |                    |                               | Amount to be<br>refunded: | \$                    |  |
|                                                                                                                                                                                                                                         |                           |                    |                               | charged                   | \$                    |  |
| a. A check in the amount of \$ to cover the above fees is enclosed.                                                                                                                                                                     |                           |                    |                               |                           |                       |  |
| A check in the amount of \$ to cover the above ites is enclosed.      Please charge my Deposit Account No. 13-4500 (Order No. 4439-4047) in the amount of \$1.332.00 to cover the above fees.                                           |                           |                    |                               |                           |                       |  |
| c. M The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to                                                                                                            |                           |                    |                               |                           |                       |  |
| Deposit Account No. 13.4500 (Order No. 4439-4047). A duplicate copy of this sheet is enclosed.  d.  Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card information should not be |                           |                    |                               |                           |                       |  |
| included on this form. Provide credit eard information and authorization on PTO-2038.                                                                                                                                                   |                           |                    |                               |                           |                       |  |
| NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the application to pending status.                                        |                           |                    |                               |                           |                       |  |
| SEND ALL CORRESPONDENCE TO:                                                                                                                                                                                                             |                           |                    |                               |                           |                       |  |
| Enter Address Here                                                                                                                                                                                                                      |                           |                    |                               |                           |                       |  |
| Morgan & Finnegan, L.L.P.                                                                                                                                                                                                               |                           |                    | - /                           |                           |                       |  |
| 3 World Financial Center Evelyn M. New York, New York 10281-2101 Name                                                                                                                                                                   |                           |                    |                               |                           |                       |  |
| Tel: (212) 415-8700 September                                                                                                                                                                                                           |                           |                    | 29, 2006                      |                           |                       |  |
| Fax: (718) 415-8701 Date 54,246                                                                                                                                                                                                         |                           |                    |                               |                           |                       |  |
| REGISTRATION NUMBER                                                                                                                                                                                                                     |                           |                    |                               |                           |                       |  |